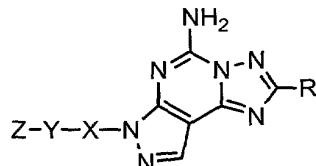


We claim:

1. Compounds having the structural formula

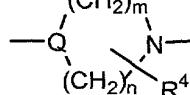


5 or a pharmaceutically acceptable salt thereof, wherein

R is R¹-furanyl, R¹-thienyl, R¹-pyridyl, R¹-pyridyl N-oxide, R¹-oxazolyl, R¹⁰-phenyl, R¹-pyrrolyl or C₄-C₆ cycloalkenyl;

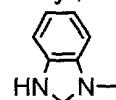
X is C₂-C₆ alkylene or -C(O)CH₂-;

Y is $-\text{N}(\text{R}^2)\text{CH}_2\text{CH}_2\text{N}(\text{R}^3)-$, $-\text{OCH}_2\text{CH}_2\text{N}(\text{R}^2)-$, $-\text{O}-$, $-\text{S}-$, $-\text{CH}_2\text{S}-$, $-(\text{CH}_2)_2\text{NH}-$, or $(\text{CH}_2)_m$



and

Z is R⁵-phenyl, R⁵-phenyl(C₁-C₆)alkyl, R⁵-heteroaryl, diphenylmethyl, R⁶-C(O)-,



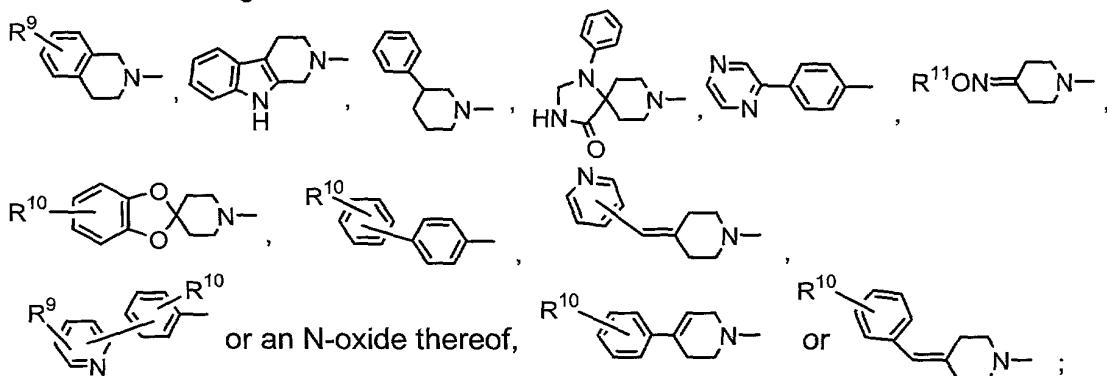
R^6-SO_2- , $R^6-OC(O)-$, $R^7-N(R^8)-C(O)-$, $R^7-N(R^8)-C(S)-$, O , phenyl- $CH(OH)-$, or



phenyl-C(=NOR²)-; or when Q is H , Z is also phenylamino or pyridylamino;

15 or

Z and Y together are



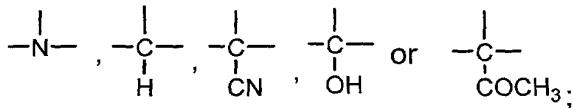
R¹ is 1 to 3 substituents independently selected from hydrogen, C₁-C₆-alkyl.

-CF₃, halogen, -NO₂, -NR¹²R¹³, C₁-C₆ alkoxy, C₁-C₆ alkylthio, C₁-C₆ alkylsulfinyl, and C₁-C₆ alkylsulfonyl;

R² and R³ are independently selected from the group consisting of hydrogen and C₁-C₆ alkyl;

m and n are independently 2-3;

Q is

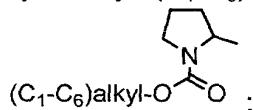


R⁴ is 1-2 substituents independently selected from the group consisting of hydrogen and C₁-C₆ alkyl, or two R⁴ substituents on the same carbon can form =O;

10 R⁵ is 1 to 5 substituents independently selected from the group consisting of hydrogen, halogen, C₁-C₆ alkyl, hydroxy, C₁-C₆ alkoxy, -CN, di-((C₁-C₆)alkyl)amino, -CF₃, -OCF₃, acetyl, -NO₂, hydroxy(C₁-C₆)alkoxy, (C₁-C₆)-alkoxy(C₁-C₆)alkoxy, di-((C₁-C₆)-alkoxy)(C₁-C₆)alkoxy, (C₁-C₆)-alkoxy(C₁-C₆)alkoxy-(C₁-C₆)-alkoxy, carboxy(C₁-C₆)-alkoxy, (C₁-C₆)-alkoxycarbonyl(C₁-C₆)alkoxy, (C₃-C₆)cycloalkyl(C₁-C₆)alkoxy, di-((C₁-C₆)alkyl)amino(C₁-C₆)alkoxy, morpholinyl, (C₁-C₆)alkyl-SO₂-, (C₁-C₆)alkyl-SO₂-(C₁-C₆)alkoxy, tetrahydropyranloxy, (C₁-C₆)alkylcarbonyl(C₁-C₆)-alkoxy, (C₁-C₆)-alkoxycarbonyl, (C₁-C₆)alkylcarbonyloxy(C₁-C₆)-alkoxy, -SO₂NH₂, phenoxy,

15
$$\begin{array}{c} (C_1-C_6 \text{ alkyl}) \\ | \\ -C=NOR^2 \end{array}, \begin{array}{c} \text{O} \\ \diagup \quad \diagdown \\ \text{O} \text{---} \text{C} \text{---} \text{CH}_3 \\ \diagdown \quad \diagup \end{array}; \text{ or adjacent } R^5 \text{ substituents together are } -O-\text{CH}_2-\text{O}-, -O-\text{CH}_2\text{CH}_2-\text{O}-, -O-\text{CF}_2-\text{O}- \text{ or } -O-\text{CF}_2\text{CF}_2-\text{O}- \text{ and form a ring with the carbon atoms to which they are attached;}$$

20 R⁶ is (C₁-C₆)alkyl, R⁵-phenyl, R⁵-phenyl(C₁-C₆)alkyl, thienyl, pyridyl, (C₃-C₆)-cycloalkyl, (C₁-C₆)alkyl-OC(O)-NH-(C₁-C₆)alkyl-, di-((C₁-C₆)alkyl)aminomethyl, or



R⁷ is (C₁-C₆)alkyl, R⁵-phenyl or R⁵-phenyl(C₁-C₆)alkyl;

25 R⁸ is hydrogen or C₁-C₆ alkyl; or R⁷ and R⁸ together are -(CH₂)_p-A-(CH₂)_q, wherein p and q are independently 2 or 3 and A is a bond, -CH₂-, -S- or -O-, and form a ring with the nitrogen to which they are attached;

R⁹ is 1-2 groups independently selected from hydrogen, C₁-C₆ alkyl, hydroxy, C₁-C₆ alkoxy, halogen, -CF₃ and (C₁-C₆)alkoxy(C₁-C₆)alkoxy;

30 R¹⁰ is 1 to 5 substituents independently selected from the group consisting of hydrogen, halogen, C₁-C₆ alkyl, hydroxy, C₁-C₆ alkoxy, -CN, -NH₂, C₁-C₆alkylamino, di-((C₁-C₆)alkyl)amino, -CF₃, -OCF₃ and -S(O)₀₋₂(C₁-C₆)alkyl;

R¹¹ is H, C₁-C₆ alkyl, phenyl, benzyl, C₂-C₆ alkenyl, C₁-C₆ alkoxy(C₁-C₆)alkyl, di-((C₁-C₆)alkyl)amino(C₁-C₆)alkyl, pyrrolidinyl(C₁-C₆)alkyl or piperidino(C₁-C₆)alkyl;

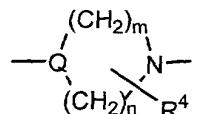
R¹² is H or C₁-C₆ alkyl; and
R¹³ is (C₁-C₆)alkyl-C(O)- or (C₁-C₆)alkyl-SO₂-.

2. A compound of claim 1 wherein R is R¹-furanyl.

5

3. A compound of claim 1 wherein X is C₂-C₆ alkylene.

4. A compound of claim 1 wherein Y is



10

5. A compound of claim 5 wherein Q is $\begin{array}{c} | \\ -N- \end{array}$ or $\begin{array}{c} | \\ -CH- \end{array}$.

6. A compound of claim 5 wherein m and n are each 2, and R⁴ is H.

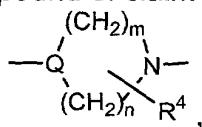
15

7. A compound of claim 1 wherein Z is R⁵-phenyl, R⁵-heteroaryl, R⁶-C(O)- or R⁶-SO₂-.

8. A compound of claim 7 wherein R⁵ is H, halogen, C₁-C₆ alkyl, C₁-C₆ alkoxy, hydroxy(C₁-C₆)alkoxy or (C₁-C₆)alkoxy(C₁-C₆)alkoxy, or R⁶ is R⁵-phenyl.

20

9. A compound of claim 1 wherein R is R¹-furanyl, X is C₂-C₆ alkylene, Y is

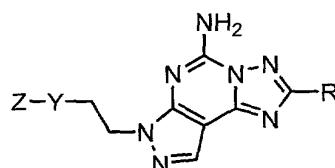


Q is $\begin{array}{c} | \\ -N- \end{array}$ or $\begin{array}{c} | \\ -CH- \end{array}$, m and n are each 2, R⁴ is H, Z is R⁵-phenyl, R⁵-heteroaryl, R⁶-C(O)- or R⁶-SO₂-.

25

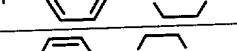
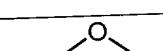
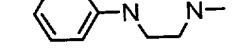
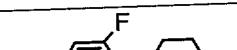
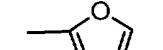
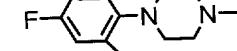
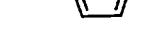
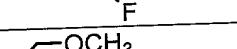
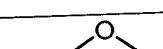
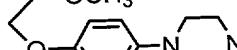
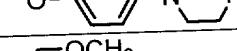
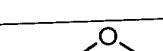
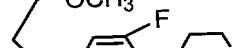
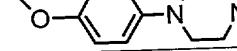
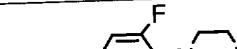
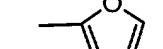
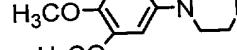
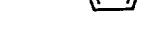
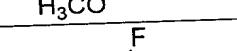
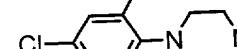
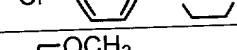
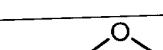
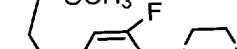
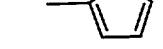
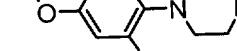
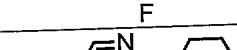
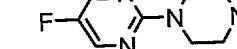
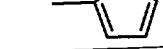
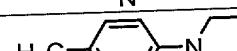
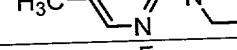
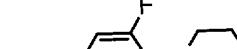
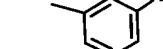
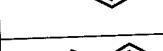
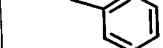
R⁵ is H, halogen, C₁-C₆ alkyl, C₁-C₆ alkoxy, hydroxy(C₁-C₆)alkoxy or (C₁-C₆)alkoxy(C₁-C₆)alkoxy, and R⁶ is R⁵-phenyl.

10. A compound of claim 1 selected from the group consisting of compounds of the formula



30

wherein R and Z-Y are as defined in the following table:

Z-Y-	R
	
	
	
	
	
	
	
	
	
	
	
	
	
	
	
	
	
	
	
	
	
	
	
	

11. A pharmaceutical composition comprising a therapeutically effective amount of a compound of claim 1 in a pharmaceutically acceptable carrier.

5 12. A method of treating central nervous system diseases or stroke, comprising
administering an effective amount of a compound of formula I to a mammal in need of
such treatment.

13. A method of claim 12 for treating depression, cognitive diseases and neurodegenerative diseases.

14. A method of claim 13 for treating Parkinson's disease, senile dementia or 5 psychoses of organic origin.

15. A process of preparing a compound of formula II



10 wherein R is R¹-furanyl, R¹-thienyl, R¹-pyridyl, R¹-pyridyl N-oxide, R¹-oxazolyl, R¹⁰-phenyl, R¹-pyrrolyl or C₄-C₆ cycloalkenyl;

15 R¹ is 1 to 3 substituents independently selected from hydrogen, C₁-C₆-alkyl, -CF₃, halogen, -NO₂, -NR¹²R¹³, C₁-C₆ alkoxy, C₁-C₆ alkylthio, C₁-C₆ alkylsulfinyl, and C₁-C₆ alkylsulfonyl;

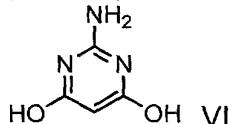
20 R¹⁰ is 1 to 5 substituents independently selected from the group consisting of hydrogen, halogen, C₁-C₆ alkyl, hydroxy, C₁-C₆ alkoxy, -CN, -NH₂, C₁-C₆alkylamino, di-((C₁-C₆)alkyl)amino, -CF₃, -OCF₃ and -S(O)₀₋₂(C₁-C₆)alkyl;

R¹² is H or C₁-C₆ alkyl; and

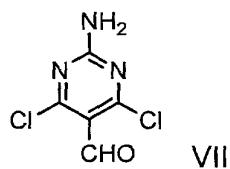
R¹³ is (C₁-C₆)alkyl-C(O)- or (C₁-C₆)alkyl-SO₂-;

comprising

25 (1) treating 2-amino-4,6-dihydroxypyrimidine

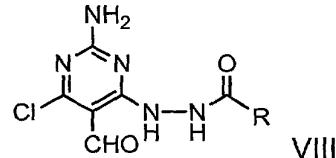


with POCl₃ in dimethylformamide to obtain 2-amino-4,6-dichloropyrimidine-5-carboxaldehyde

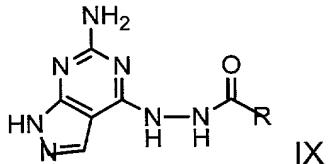


25 (2) treating carboxaldehyde VII with a hydrazide of the formula

H₂N-NH-C(O)-R, wherein R is as defined above, to obtain



(3) treating the intermediate of formula VIII with hydrazine hydrate to form a pyrazolo ring, thus obtaining the intermediate of formula IX

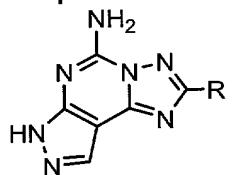


IX

(4) forming the desired compound of formula II by dehydrative rearrangement.

5

16. A process for preparing a compound of the formula II



II

wherein R is R¹-furanyl, R¹-thienyl, R¹-pyridyl, R¹-pyridyl N-oxide, R¹-oxazolyl, R¹⁰-phenyl, R¹-pyrrolyl or C₄-C₆ cycloalkenyl;

10 R¹ is 1 to 3 substituents independently selected from hydrogen, C₁-C₆-alkyl, -CF₃, halogen, -NO₂, -NR¹²R¹³, C₁-C₆ alkoxy, C₁-C₆ alkylthio, C₁-C₆ alkylsulfinyl, and C₁-C₆ alkylsulfonyl;

15 R¹⁰ is 1 to 5 substituents independently selected from the group consisting of hydrogen, halogen, C₁-C₆ alkyl, hydroxy, C₁-C₆ alkoxy, -CN, -NH₂, C₁-C₆ alkylamino, di-((C₁-C₆)alkyl)amino, -CF₃, -OCF₃ and -S(O)₀₋₂(C₁-C₆)alkyl;

R¹² is H or C₁-C₆ alkyl; and

R¹³ is (C₁-C₆)alkyl-C(O)- or (C₁-C₆)alkyl-SO₂-;

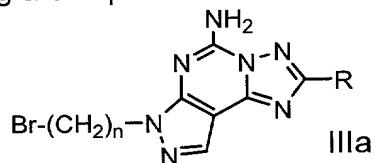
comprising converting a compound of formula IX



IX

20 into the desired compound of formula II by dehydrative rearrangement.

17. A process for preparing a compound of formula IIIa



IIIa

wherein R is R¹-furanyl, R¹-thienyl, R¹-pyridyl, R¹-pyridyl N-oxide, R¹-oxazolyl,

25 R¹⁰-phenyl, R¹-pyrrolyl or C₄-C₆ cycloalkenyl;

R¹ is 1 to 3 substituents independently selected from hydrogen, C₁-C₆-alkyl, -CF₃, halogen, -NO₂, -NR¹²R¹³, C₁-C₆ alkoxy, C₁-C₆ alkylthio, C₁-C₆ alkylsulfinyl, and C₁-C₆ alkylsulfonyl;

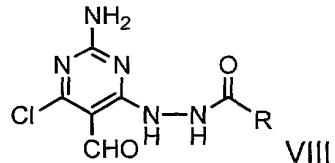
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R¹⁰ is 1 to 5 substituents independently selected from the group consisting of hydrogen, halogen, C₁-C₆ alkyl, hydroxy, C₁-C₆ alkoxy, -CN, -NH₂, C₁-C₆alkylamino, di-((C₁-C₆)alkyl)amino, -CF₃, -OCF₃ and -S(O)₀₋₂(C₁-C₆)alkyl;

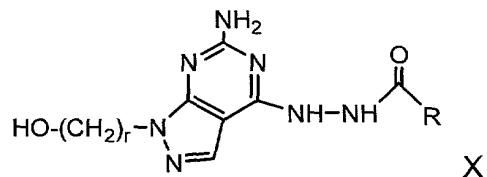
R¹² is H or C₁-C₆ alkyl; and

5 R¹³ is (C₁-C₆)alkyl-C(O)- or (C₁-C₆)alkyl-SO₂-; comprising

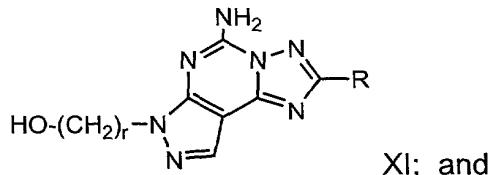
(1) treating a chloride of formula VIII



10 with a hydroxyalkyl hydrazine of the formula HO-(CH₂)_r-NNHH₂, wherein r is 2-6, to obtain



(2) cyclizing the intermediate of formula X by dehydrative rearrangement to obtain the tricyclic intermediate of formula XI



15 (3) converting the hydroxy compound of formula XI to the bromide of formula IIIa.

18. A pharmaceutical composition comprising a therapeutically effective amount of a combination of a compound of claim 1 and 1 to 3 other agents useful in treating

20 Parkinson's disease in a pharmaceutically acceptable carrier

19. A method of treating Parkinson's disease comprising administering to a mammal in need of such treatment an effective amount of a combination of a compound of claim 1 and 1 to 3 other agents useful in treating Parkinson's disease.

25 20. The method of claim 19 wherein the other agents are selected from the group consisting of L-DOPA, dopaminergic agonists, MAO-B inhibitors, DOPA decarboxylase inhibitors and COMT inhibitors.